Application No.: 10/058,215

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

1. (Currently Amended) A compound having the Formula IV:

wherein:

R¹, R², R³, R⁴ and R⁵ independently represent hydrogen, halogen, alkyl, aryl, aralkyl, heteroaryl or heteroarylalkyl;

R⁶, R⁷, R⁸ and R⁹ independently represent hydrogen, alkyl, hydroxyalkyl, aminoalkyl, monoalkylaminoalkyl, dialkylaminoalkyl, carboxyalkyl, aryl or aralkyl;

or R^6 and R^7 are taken together to form -(CH₂)_p-, where p is 2-8, while R^8 and R^9 are defined as above; or R^8 and R^9 are taken together to form -(CH₂)_q-, where q is 2-8, while R^6 and R^7 are defined as above; or R^6 and R^8 are taken together to form -(CH₂)_r-, while r is zero (a bond), 1 or 2, while R^7 and R^9 are defined as above;

X represents oxygen, sulfur, $-CH_2$ -, -NH-, -(C=O)NH- or -NH(C=O)-; n is from 0 to 4; m is from 0 to 4;

a is 0 or 1;

D represents oxygen;

v is 0 or 1;

R¹⁰, R¹¹, R¹² and R¹³ independently represent: hydrogen; hydroxy; alkyl; alkoxy; cycloalkyl; aryl, optionally substituted with one or more of halogen, hydroxy, cyano, alkyl, aryl, alkoxy, haloalkyl, arylalkoxy, aryloxy, alkylsulfonyl, alkylsulfinyl, alkylalkoxyaryl, monoalkylamino, dialkylamino, aminoalkyl, monoalkylaminoalkyl, dialkylaminoalkyl, alkanoyl; monoalkylamino; dialkylamino; aminoalkyl;

Application No.: 10/058,215

monoalkylaminoalkyl; dialkylaminoalkyl; alkanoyl; heteroaryl having 5-14 ring members, optionally substituted with one or more of halogen, hydroxy, cyano, alkyl, aryl, alkoxy, haloalkyl, arylalkoxy, aryloxy, alkylsulfonyl, alkylsulfinyl, alkylalkoxyaryl, monoalkylamino, dialkylamino, aminoalkyl, monoalkylaminoalkyl, dialkylaminoalkyl, alkanoyl; or

wherein $\ensuremath{R^{17}}$ and $\ensuremath{R^{18}}$ together form -CH2CH2-O-, -O-CH2CH2-,

-O-CH₂-O- or -O-CH₂CH₂-O-; or

 R^{10} and R^{12} are taken together to form -(CH₂)_s-, wherein s is 0 (a bond) or 1 to 4, while R^{11} and R^{13} are as defined as above; or R^{10} and R^{12} are taken together to form a double bond when i is 0 and k is 1, while R^{11} and R^{13} are as defined above; or R^{10} and R^{11} are taken together to form -(CH₂)_t-, wherein t is 2 to 8, while R^{12} and R^{13} are as defined as above, or R^{12} and R^{13} are taken together to form -(CH₂)_u- wherein u is 2 to 8, while R^{10} and R^{11} are as defined as above;

i is from 0 to 4;

j is from 0 to 4;

k is 0 or 1;

R¹⁴ is hydrogen or a functionality that acts as a prodrug, including alkyl, aryl, aralkyl, dialkylaminoalkyl, 1-morpholinoalkyl, 1-piperidinylalkyl, pyridinylalkyl, alkoxy(alkoxy) alkoxyalkyl, or (alkoxycarbonyl)oxyethyl;

W is:

Application No.: 10/058,215

$$\frac{1}{1} \frac{1}{1} \frac{1}$$

wherein:

Y is N or CH;

Z is N- or -CH;

R¹⁵-is hydrogen, halogen, alkyl, aryl or arylalkyl;

R¹⁶ is hydrogen, alkyl, haloalkyl or halogen; and

R¹⁹ and R²⁰ are independently hydrogen, halogen or alkyl;

 R^{27} , R^{28} , and R^{29} , R^{30} and R^{31} are independently hydrogen, halogen, alkyl, alkoxy or aryl; and o and p are independently 0, 1 or 2.

- 2. (Original) The compound of claim 1, wherein R¹⁴ is a prodrug, selected from the group consisting of: alkyl, aryl, aralkyl, dialkylaminoalkyl, 1-morpholinoalkyl, 1-piperidinylalkyl, pyridinylalkyl, alkoxy(alkoxy) alkoxyalkyl, or (alkoxycarbonyl)oxyethyl.
- 3. (Currently Amended) The compound of claim 1, wherein:

 R^1 , R^2 , R^3 , R^4 and R^5 independently represent hydrogen, halogen, (C_{1-8}) alkyl, (C_{6-10}) aryl, (C_{6-10}) ar (C_{1-8}) alkyl, 5-14 member heteroaryl, or 5-14 member heteroaryl (C_{1-8}) alkyl;

 R^6 , R^7 , R^8 and R^9 independently represent hydrogen, (C_{1-8}) alkyl, hydroxy (C_{1-8}) alkyl, amino (C_{1-8}) alkyl, mono (C_{1-8}) alkylamino (C_{1-8}) alkyl, di (C_{1-8}) alkylamino (C_{1-8}) alkyl, carboxy (C_{1-8}) alkyl, (C_{6-10}) aryl or (C_{6-10}) ar (C_{1-8}) alkyl;

Application No.: 10/058,215

or R^6 and R^7 are taken together to form - $(CH_2)_p$ -, where p is 2-8, while R^8 and R^9 are defined as above; or R^8 and R^9 are taken together to form - $(CH_2)_q$ -, where q is 2-8, while R^6 and R^7 are defined as above; or R^6 and R^8 are taken together to form - $(CH_2)_r$ -, while r is zero (a bond), 1 or 2, while R^7 and R^9 are defined as above;

X represents oxygen, sulfur, -CH₂-, -NH-, -(C=O)NH- or -NH(C=O)-; n is from 0 to 4; m is from 0 to 4; a is from 0 or 1; D represents oxygen;

v is from 0 or 1;

R¹⁰, R¹¹, R¹² and R¹³ independently represent: hydrogen; hydroxy; (C₁₋₈)alkyl; (C₁₋₈)alkoxy; (C₃₋₈)cycloalkyl; (C₆₋₁₀)aryl, optionally substituted with one or more of halogen, hydroxy, cyano, (C₁₋₈)alkyl, (C₆₋₁₀)aryl, (C₁₋₈)alkoxy, halo(C₁₋₈)alkyl, (C₆₋₁₀)aryl(C₁₋₈)alkyl, (C₆₋₁₀)aryl(C₁₋₈)alkyl, (C₆₋₁₀)aryl(C₁₋₈)alkylsulfonyl, (C₁₋₈)alkylsulfinyl, (C₁₋₈)alkylsulfinyl, (C₁₋₈)alkoxy(C₆₋₁₀)aryl(C₁₋₈)alkyl, mono(C₁₋₈)alkylamino, di(C₁₋₈)alkylamino, amino(C₁₋₈)alkyl, mono(C₁₋₈)alkylamino; di(C₁₋₈)alkylamino; amino(C₁₋₈)alkyl; mono(C₁₋₈)alkyl, (C₁₋₈)alkyl; di(C₁₋₈)alkylamino(C₁₋₈)alkyl; (C₁₋₈)alkyl; mono(C₁₋₈)alkyl; di(C₁₋₈)alkylamino(C₁₋₈)alkyl; (C₁₋₈)alkanoyl; heteroaryl having 5-14 ring members, optionally substituted with one or more of halogen, hydroxy, cyano, (C₁₋₈)alkyl, (C₆₋₁₀)aryl, (C₁₋₈)alkoxy, halo(C₁₋₈)alkyl, (C₆₋₁₀)aryl(C₁₋₈)alkyl, (C₆₋₁₀)aryl(C₁₋₈)alkoxy, (C₆₋₁₀)aryloxy, (C₁₋₈)alkylsulfonyl, (C₁₋₈)alkylsulfinyl, (C₁₋₈)alkoxy(C₆₋₁₀)aryl(C₁₋₈)alkyl, mono(C₁₋₈)alkylamino, di(C₁₋₈)alkylamino, amino(C₁₋₈)alkyl, mono(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, (C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, (C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, (C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkyl, di(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)alkylamino(C₁₋₈)

wherein R^{17} and R^{18} together form -CH₂CH₂-O-, -O-CH₂CH₂-, -O-CH₂CH₂-O- or -O-CH₂CH₂-O-; or

Application No.: 10/058,215

 R^{10} and R^{12} are taken together to form -(CH₂)_s-, wherein s is 0 (a bond) or 1 to 4, while R^{11} and R^{13} are as defined as above; or R^{10} and R^{12} are taken together to form a double bond when i is 0 and k is 1, while R^{11} and R^{13} are as defined above; or R^{10} and R^{11} are taken together to form -(CH₂)_t-, wherein t is 2 to 8, while R^{12} and R^{13} are as defined as above, or R^{12} and R^{13} are taken together to form -(CH₂)_u- wherein u is 2 to 8, while R^{10} and R^{11} are as defined as above; or

i is from 0 to 4;

j is from 0 to 4; and

k is 0 or 1; `..

R¹⁴ is hydrogen or a functionality that acts as a prodrug;

W is:

wherein:

Y is N or CH;

Z is N or CH;

 $R^{15} \hbox{-is hydrogen, halogen, } (C_{1-8}) \hbox{alkyl, } (C_{6-10}) \hbox{aryl or } (C_{6-10}) \hbox{aryl(} C_{1-8}) \hbox{alkyl;}$

DOCKET NO.: 3DP-0418 (1030002) PATENT

Application No.: 10/058,215

 R^{16} is hydrogen, (C_{1-8}) alkyl, halo (C_{1-8}) alkyl or halogen; R^{19} -and R^{20} -are independently hydrogen, halogen or (C_{1-8}) alkyl; and R^{27} , R^{28} , and R^{29} , R^{30} -and R^{31} are independently hydrogen, halogen, (C_{1-8}) alkyl, (C_{1-8}) alkoxy, or (C_{6-10}) aryl-and o and p are independently 0, 1 or 2.

- 4. (Original) The compound of claim 1, wherein R^1 and R^2 independently represent hydrogen, halogen, (C_{1-6}) alkyl, (C_{6-10}) aryl, (C_{6-10}) aryl, (C_{1-6}) alkyl, 5-14 member heteroaryl, or 5-14 member heteroaryl (C_{1-8}) alkyl.
- 5. (Original) The compound of claim 4, wherein R¹ and R² independently represent hydrogen, methyl, ethyl, propyl, butyl, phenyl, benzyl or phenylethyl.
- 6. (Original) The compound of claim 5, wherein R¹ and R² independently represent hydrogen, methyl, ethyl or propyl.
- 7. (Original) The compound of claim 1, wherein R^3 , R^4 and R^5 independently represent hydrogen, (C_{1-6}) alkyl, (C_{6-10}) aryl, or (C_{6-10}) aryl, or (C_{1-6}) alkyl.
- 8. (Original) The compound of claim 7, wherein R^3 , R^4 and R^5 are hydrogen or (C_{1-4}) alkyl.
- 9. (Original) The compound of claim 1, wherein R^6 , R^7 , R^8 and R^9 independently represent hydrogen, halogen or (C_{1-6})alkyl.
- 10. (Original) The compound of claim 1, wherein X is oxygen, -CH₂- or -(C=O)NH-.
- 11. (Original) The compound of claim 10, wherein X is oxygen or -CH₂-.
- 12. (Currently Amended) The compound of claim 1, wherein W is:

wherein:

Y-is-N-or-CH;

 $R^{15} \text{-is hydrogen, halogen, } (C_{1-8}) \\ alkyl, \\ (C_{6-10}) \\ aryl \\ \text{or } (C_{6-10}) \\ aryl(C_{1-8}) \\ alkyl; \\$

 R^{16} is hydrogen, (C_{1-8}) alkyl, halo (C_{1-8}) alkyl or halogen; and

R¹⁹-and R²⁰-are hydrogen, halogen or (C₁₋₈)alkyl; and

 R^{27} , R^{28} , and R^{29} , R^{30} and R^{31} are hydrogen, halogen, (C_{1-8}) alkyl, (C_{1-8}) alkoxy, or (C_{6-10}) aryl.

13. (Currently Amended) The compound of claim 1, wherein W is

wherein:

 R^{15} -is hydrogen, halogen, (C_{1-8}) alkyl, (C_{6-10}) aryl or (C_{6-10}) aryl (C_{1-8}) alkyl;

PATENT

DOCKET NO.: 3DP-0418 (1030002)

Application No.: 10/058,215

 R^{16} is hydrogen, (C_{1-8}) alkyl, halo (C_{1-8}) alkyl or halogen; <u>and</u> R^{19} -and R^{20} -are hydrogen, halogen or (C_{1-8}) alkyl; and R^{27} , R^{28} , <u>and</u> R^{29} , R^{30} -and R^{31} are hydrogen, halogen, (C_{1-8}) alkyl, (C_{1-8}) alkoxy, or (C_{6-10}) aryl.

14. (Currently Amended) The compound of claim 13, wherein W is

- 15. (Original) The compound of claim 1, wherein R^{10} , R^{11} , R^{12} and R^{13} independently represent: hydrogen; hydroxy, (C_{1-12}) alkyl; (C_{3-6}) cycloalkyl; or (C_{6-10}) aryl, optionally substituted with one or more of halogen, hydroxy, cyano, (C_{1-8}) alkyl, (C_{6-10}) aryl, (C_{1-8}) alkoxy, halo (C_{1-8}) alkyl, (C_{6-10}) aryl (C_{1-8}) alkyl, (C_{6-10}) aryloxy.
- 16. (Original) The compound of claim 1, wherein R^{10} , R^{11} , R^{12} and R^{13} independently represent a heteroaryl having 5-14 ring members, optionally substituted with one or more of halogen, hydroxy, cyano, (C_{1-8}) alkyl, (C_{1-8}) alkoxy (C_{6-10}) aryl, (C_{1-8}) alkoxy, halo (C_{1-8}) alkyl, (C_{6-10}) aryl (C_{1-8}) alkyl, (C_{6-10}) aryl (C_{1-8}) alkyl, (C_{6-10}) aryl (C_{1-8}) alkylamino, (C_{1-8}) alkylamino, di (C_{1-8}) alkylamino, di (C_{1-8}) alkylamino, or carboxy (C_{1-8}) alkyl; or

wherein R^{17} and R^{18} together form -CH₂CH₂-O-, -O-CH₂CH₂-, -O-CH₂-O- or -O-CH₂CH₂-O-.

17. (Original) The compound of claim 1, wherein R^{12} and R^{13} are independently thiazolyl, benzofuranyl,

Application No.: 10/058,215

$$\mathbb{R}^{34}$$
 or \mathbb{R}^{33}

wherein:

b is from 0 to 4;

 R^{32} is halogen, (C_{1-6}) alkyl, halo (C_{1-6}) alkyl, (C_{1-6}) alkoxy,

 (C_{1-6}) alkoxy (C_{1-6}) alkyl or halo (C_{1-6}) alkoxy;

R³³ is halogen; and

 R^{34} is (C_{1-6}) alkyl, hydroxy or (C_{1-6}) alkoxy, or

two of R³², or two of R³³, or one of R³³ and R³⁴, when attached to adjacent carbon atoms, may together form a ring, wherein the ring formed is an aliphatic, aryl or heteroaryl ring, each of which may be optionally substituted by one or more of halogen, hydroxy, cyano, alkyl, aryl, alkoxy, haloalkyl, arylalkyl, arylalkoxy, aryloxy, alkylsulfonyl, alkylsulfinyl, alkoxyarylalkyl, monoalkylamino, dialkylamino, aminoalkyl, monoalkylaminoalkyl, dialkylamino; dialkylamino; aminoalkyl; monoalkylaminoalkyl; dialkylaminoalkyl; alkanoyl.

- 18. (Original) The compound of claim 1, wherein R^{10} and R^{12} are taken together to form a double bond where i is 0 and k is 1, and R^{11} and R^{13} are each hydrogen.
- 19. (Original) The compound of claim 1, wherein R^{10} is an optionally substituted aryl or an optionally substituted heteroaryl.
- 20. (Original) The compound of claim 1, wherein i and j are each 0.
- 21. (Original) The compound of claim 1, wherein k is 1.
- 22. (Original) The compound of claim 1, wherein R^{14} is hydrogen.

PATENT

DOCKET NO.: 3DP-0418 (1030002)

Application No.: 10/058,215

23. (Original) The compound of claim 1, wherein i and j are each zero; k is one; R^{10} , R^{11} and R^{12} are each hydrogen; and R^{13} is hydrogen, (C_{1-6}) alkyl, (C_{6-10}) aryl or (C_{6-10}) ar(C_{1-4})alkyl.

24. (Currently Amended) The compound of claim 1, wherein:

 R^1 is hydrogen or (C_{1-4}) alkyl;

 R^2 , R^3 , R^4 , and R^5 are hydrogen or (C_{1-4})alkyl;

 R^6 , R^7 , R^8 and R^9 are hydrogen or (C_{1-4}) alkyl;

X is oxygen or -CH₂-;

n is 0 or 1;

m is 0 or 1;

 R^{10} , R^{11} , R^{12} and R^{13} independently represent hydrogen, (C_{1-6}) alkyl or (C_{6-10}) ar (C_{1-6}) alkyl; or

one of the combination R¹⁰ and R¹¹, R¹² and R¹³ or R¹⁰ and R¹² are taken together to form -(CH₂)_s-, wherein s is 1 or 2 while the remaining of R¹⁰-R¹³ are as defined above;

i is 0 or 1;

j is 0 or 1;

k is 0 or 1;

R¹⁴ is hydrogen, C₁₋₆ alkyl or benzyl;

W is:

wherein:

Y is N or CH;

 R^{15} is hydrogen, halogen, (C_{1-8}) alkyl, (C_{6-10}) aryl or (C_{6-10}) aryl (C_{1-8}) alkyl;

PATENT

DOCKET NO.: 3DP-0418 (1030002)

Application No.: 10/058,215

 $R^{16} \text{ is hydrogen, } (C_{1\text{-8}}) \text{alkyl, halo} (C_{1\text{-8}}) \text{alkyl or halogen; } \underline{\text{and}}$ $R^{19} \text{-and } R^{20} \text{-are hydrogen, halogen or } (C_{1\text{-8}}) \text{alkyl; and}$ $R^{27}, R^{28}, \underline{\text{and}} R^{29}, R^{30} \text{-and } R^{31} \text{ are hydrogen, halogen, } (C_{1\text{-8}}) \text{alkyl, } (C_{1\text{-8}}) \text{alkoxy, or } (C_{6\text{-10}}) \text{aryl.}$

- 25. (Original) The compound of claim 24, wherein R^1 , R^2 , R^3 , R^4 , R^5 , R^6 , R^7 , R^8 , R^9 , R^{11} , R^{12} , R^{13} and R^{14} are hydrogen.
- 26. (Original) The compound of claim 1, wherein R^{10} and R^{11} are taken together to form $(CH_2)_t$ -, where t is 2 to 5 and R^{12} and R^{13} are each hydrogen.
- 27. (Original) The compound of claim 1, wherein R^{12} and R^{13} are taken together to form $(CH_2)_u$ -, where u is 2 to 5 and R^{10} and R^{11} are each hydrogen.
- 28. (Original) The compound of claim 1, wherein R^{10} and R^{12} are taken together to form $(CH_2)_s$ where s is zero or 1 to 4, and R^{11} and R^{13} are each hydrogen.
- 29. (Original) The compound of claim 1, wherein:

X is -(C=O)NH-;

n, m, a and v are each 0; and

 R^6 , R^7 , R^{12} and R^{13} are hydrogen.

30. (Original) The compound of claim 1, wherein:

X is oxygen;

n and m are each 0;

a and v are each 1;

D is oxygen;

R⁶, R⁷, R⁸ and R⁹ are hydrogen.

31. (Original) The compound of claim 1, wherein:

X is oxygen;

n, m and v are each 0;

DOCKET NO.: 3DP-0418 (1030002) **Application No.:** 10/058,215

a is 1; and R^6 , R^7 , R^{12} and R^{13} are hydrogen.

32. (Original) The compound of claim 1, wherein:

X is -CH₂-;

n, m and v are each 0;

a is 1; and

R⁶, R⁷, R¹² and R¹³ are hydrogen.

- 33. (Original) The compound of claim 1, wherein v is 0.
- 34. (Currently Amended) The compound of claim 1, wherein

R¹ is hydrogen or -CH₃;

 R^2 , R^3 , R^4 , R^5 , R^6 , R^7 , R^8 and R^9 are hydrogen;

X is oxygen or $-CH_2$ -;

n is 0 or 1;

m is 0 or 1;

R¹⁰, R¹¹, R¹² and R¹³ independently represent hydrogen,

 (C_1-C_6) alkyl or (C_{6-10}) ar (C_{1-6}) alkyl; or

one of the combination R¹⁰ and R¹¹, R¹² and R¹³ or R¹⁰ and R¹² are taken together to form -(CH₂)_s-, wherein s is 1 while the remaining of R¹⁰-R¹³ are defined above;

i is 0 or 1;

j is 0 or 1;

k is 0 or 1;

R¹⁴ is hydrogen or alkyl;

W is:

01

Application No.: 10/058,215

wherein:

 R^{15} is hydrogen, halogen, (C_{1-8}) alkyl, (C_{6-10}) aryl or (C_{6-10}) aryl (C_{1-8}) alkyl;

R¹⁶ is hydrogen, (C₁₋₈)alkyl, halo(C₁₋₈)alkyl or halogen; and

 R^{27} , R^{28} , and R^{29} , R^{30} and R^{31} are hydrogen, halogen, (C_{1-8}) alkyl, (C_{1-8}) alkoxy, or (C_{6-10}) aryl.

35. (Original) The compound of claim 1, wherein:

v, m, n, i and j are 0;

a is 1;

 R^1 , R^2 , R^3 , R^4 , R^5 , R^6 , R^7 , R^8 , R^9 , R^{11} , R^{12} , R^{13} and R^{14} are each hydrogen; and R^{10} is pyridinyl.

36. (Original) The compound of claim 1, wherein:

v, m, n, i and j are 0;

a is 1;

 R^1 , R^2 , R^3 , R^4 , R^5 , R^6 , R^7 , R^8 , R^9 , R^{11} , R^{12} , R^{13} and R^{14} are each hydrogen; and R^{10} is quinolinyl.

37. (Original) The compound of claim 1, wherein:

v, m, n, i and j are 0;

a is 1;

 R^1 , R^2 , R^3 , R^4 , R^5 , R^6 , R^7 , R^8 , R^9 , R^{11} , R^{12} , R^{13} and R^{14} are each hydrogen; and R^{10} is methoxyphenyl.

38. (Currently Amended) The compound of claim 1, which is one of:

3-{5-[3-(2-pyridylamino)propoxy]indolyl}propanoic acid;

-----3-{5-[3-(2-pyridylamino)propoxy]indolyl}acetic acid;

---- 3-{2-methyl-5-[3-(2-pyridylamino)propoxy]indolyl}propanoic acid;

Application No.: 10/058,215

- 2 (trans 2 {5-[3-(2-pyridylamino)propoxy]indolyl} cyclopropyl)acetic acid;
- -----3-(5-{2-[6 (methylamino)-2-pyridyl]ethoxy} indolyl)propanoic acid;
- -----2-benzyl 3-{5[3 (2-pyridylamino)propoxy]indolyl}propanoic acid;
- 2-methyl-3-{5-[3-(2-pyridylamino)propoxy]indolyl}propanoic acid;
- 2 ({5-[3-(2-pyridylamino)propoxy]indolyl}methyl)pentanoic acid;
- 2 ({5-[3-(2 pyridylamino)propoxy]indolyl}methyl)octanoic acid;
- 3-[5-(3-{[benzylamino]carbonylamino}propoxy)indolyl] propanoic acid;
- 3-[5-(2-5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl-acetylamino)-indol-1-yl]-hexanoic acid;
- 3 (5 {2 [N (4,5 dihydro-1H imidazol 2-yl)-aminooxy]-ethoxy}-indol-1 yl) 3 phenyl-propionic acid;
- 3 (5 {2 [guanidino-oxy]-ethoxy}-indol-1-yl)-3-phenyl-propionic acid;
- 3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-hexanoic acid;
- 3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-phenyl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-phenyl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(3-benzyloxy-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-3-p-tolyl-propionic acid;
- 3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-3-m-tolyl-propionic acid;
- 3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-3-o-tolyl-propionic acid;
- 3-biphenyl-4-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;

Application No.: 10/058,215

- 3-(3,5-dichloro-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(3,5-difluoro-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(3-cyano-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(4-cyano-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(2-methoxy-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(3-methoxy-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(4-methoxy-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-quinolin-3-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(3-chloro-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-naphthalen-2-yl-3-{5-[2-(5,6,7,8-Tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(2-chloro-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-naphthalen-1-yl-3-{5-[2-(5,6,7,8-Tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-(4-fluoro-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-3-(3-trifluoromethyl-phenyl)-propionic acid;
- 3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-3-(4-trifluoromethyl-phenyl)-propionic acid;

Application No.: 10/058,215

3-pyridin-3-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;

3-pyridin-2-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;

3-pyridin-4-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-acrylic acid;

3-(2,3-dihydro-benzofuran-5-yl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;

3-benzo[1,3]dioxol-5-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;

3-(5-methanesulfonyl-pyridin-3-yl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethyoxy]-indol-1-yl}-propionic acid;

- 3-{5-[2-(6-methylamino-pyridin-2-yl)-ethoxy]-indol-1-yl}-3-phenyl-propionic acid;
- 3-{5-[2-(6-methylamino-pyridin-2-yl)-ethoxy]-indol-1-yl}-3-quinolin-3-yl-propionic acid;
- 3-{5-[2-(6-methylamino-pyridin-2-yl) ethoxy] indol-1-yl}-3-pyridin-3-yl-propionic acid;
- 3 {5 [2 (6 methylamino-pyridin-2-yl) ethoxy] indol-1-yl} hexanoic acid;
- 3-{5-[2-(2-methyl-5,6,7,8-tetrahydro-[1,8]naphthyridin-3-yl)-ethyl]-indol-1-yl}-propionic acid;
- 3-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-indol-1-yl}-propionic acid;
- 3-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-indol-1-yl}-hexanoic acid;
- 3-phenyl-3-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-indol-1-yl}-propionic acid;
- 3 {5 [2 (3,4 dihydro 2H pyrido[3,2 b][1,4]oxazin 6 yl) ethoxy] indol 1 yl} 3 phenyl propionic acid;
- 3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-3-[5-(2,2,2-trifluoro-ethoxy)-pyridin-3-yl]-propionic acid;

Application No.: 10/058,215

3-(5-ethoxy-pyridin-3-yl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;

3-pyridin-4-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;

3-pyridin-2-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-acrylic acid;

6 (2 hydroxy ethyl) 2,3 dihydro-pyrido[3,2 b][1,4]oxazine 4 carboxylic acid tert-butyl ester;

3 {5 [2 (6 methylamino pyridin 2 yl) ethoxy] indol-1 yl} 3 quinolin 3 yl-propionic acid;

or a pharmaceutically acceptable salt, hydrate, solvate or prodrug thereof.

- 39. (Currently Amended) The compound of claim 1, which is one of:
- 3-(3-methoxy-phenyl)-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-quinolin-3-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-pyridin-3-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3-pyridin-2-yl-3-{5-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-indol-1-yl}-propionic acid;
- 3 {5 [2 (6 methylamino pyridin 2 yl) ethoxy] indol 1 yl} 3 quinolin 3 yl propionic acid;
- or a pharmaceutically acceptable salt, hydrate, solvate or prodrug thereof.
- 40. (Original) A pharmaceutical composition comprising a compound of claim 1 and a pharmaceutically acceptable carrier or diluent.
- 41. (Previously Presented) A method of treating a pathological condition selected from the group consisting of osteoporosis, macular degeneration, diabetic retinopathy, rheumatoid

Application No.: 10/058,215

arthritis and sickle cell anemia, in a mammal in need of such treatment, comprising administering to said mammal an effective amount of a compound of claim 1.

- 42. (Canceled)
- 43. (Original) The method of claim 41, wherein said condition is osteoporosis.
- 44. (Canceled)
- 45. (Canceled)
- 46. (Original) The method of claim 41, wherein said condition is macular degeneration.
- 47. (Original) The method of claim 41, wherein said condition is diabetic retinopathy.
- 48. (Original) The method of claim 41, wherein said condition is rheumatoid arthritis.
- 49. (Original) The method of claim 41, wherein said condition is sickle cell anemia.
- 50. (Original) A process for preparing a substituted indole compound of claim 1, comprising:

reacting a compound of Formula V:

$$R^{5}$$
 R^{2}
 R^{1}
 R^{10}
 R^{11}
 R^{12}
 R^{13}
 R^{13}
 R^{14}
 R^{14}
 R^{15}
 R^{15}

or a salt, hydrate or solvate thereof, wherein R¹, R², R³, R⁴, R⁵, R¹⁰, R¹¹, R¹², R¹³, R¹⁴, i and j are as defined in claim 1,

Application No.: 10/058,215

with the compound of Formula VI or Formula X:

$$R^{15}$$

OH

Or

 R^{27}
 R^{28}
 R^{16}
 R^{16}
 R^{16}

or a salt, hydrate or solvate thereof, wherein R¹⁵, R¹⁶, R²⁷, R²⁸ and R²⁹ are as defined in claim 1, to form a substituted indole compound of claim 1.

Claims 51-53 (Canceled)

- 54. (Original) A method for treating a central nervous system (CNS) related disorder, selected from the group consisting of: neuronal loss associated with stroke, ischemia, CNS trauma, hypoglycemia, surgery, a neurodegenerative disease, an adverse consequence of overstimulation of one or more excitatory amino acids, schizophrenia, anxiety, convulsions, chronic pain, psychosis, anesthesia, and opiate tolerance, in a mammal in need of such treatment, comprising administering to said mammal an effective amount of a compound of claim 1.
- 55. (Original) The method according to claim 54, wherein said CNS related disorder is neuronal loss associated with stroke.
- 56. (Original) The method according to claim 54, wherein said CNS related disorder is ischemia.
- 57. (Original) The method according to claim 54, wherein said CNS related disorder is CNS trauma.
- 58. (Original) The method according to claim 54, wherein said CNS related disorder is hypoglycemia.

59. (Original) The method according to claim 54, wherein said CNS related disorder is the

result of surgery.

60. (Original) The method according to claim 54, wherein said CNS related disorder is a

neurodegenerative disease.

61. (Original) The method according to claim 60, wherein said neurodegenerative disease is

selected from Alzheimer's disease or Parkinson's disease.

62. (Original) The method according to claim 54, wherein said CNS related disorder is

schizophrenia.

63. (Original) The method according to claim 41, wherein the activity of $\alpha 4$ integrin is

inhibited.

64. (Original) The method according to claim 54, wherein the activity of α 4 integrin is

inhibited.

65. (Original) The compound according to claim 17, wherein R^{12} and R^{13} are independently

selected from:

\$-